

AMENDMENTS TO THE CLAIMS

1. (withdrawn) A method of identifying a condition in an individual in which an elevated serum or plasma HDL concentration, or HDL cholesterol concentration, provides enhanced protection against cardiovascular disease, the method comprising the step of testing the individual for a disorder that detrimentally affects the protective effect of HDL, whereby absence of such a disorder is an indication of enhanced protection against cardiovascular disease when said individual exhibits elevated serum or plasma HDL or HDL cholesterol concentration.

2. (withdrawn) The method according to claim 1, wherein the disorder is selected from liver damage and oxidative stress.

3. (withdrawn) The method according to claim 1 or 2, comprising the additional step of determining the serum or plasma HDL or HDL cholesterol concentration.

4. (withdrawn) The method according to claim 2 wherein the disorder comprises liver damage and testing for liver damage comprises determining  $\gamma$ -glutamyltransferase or a liver transaminase activity or concentration in a serum or plasma sample and comparing it to a selected reference value for  $\gamma$ -glutamyltransferase.

5. (withdrawn) The method according to claim 2, wherein the disorder comprises liver damage and testing for liver damage comprises genotyping mutations or polymorphisms inducing or predisposing to liver damage or influencing serum or plasma  $\gamma$ -glutamyltransferase activity or concentration or the testing of the expression of the genes encoding these proteins.

6. (withdrawn) The method according to claim 2, wherein the disorder comprises oxidative stress and testing for oxidative stress comprises determining serum or plasma activity or concentration of one or several phase I or phase II detoxification enzyme.

7. (withdrawn) The method according to claim 2, wherein the disorder comprises oxidative stress and testing of oxidative stress is the assessment of serum or plasma concentration of ferritin or an oxidized fatty acid, oxidized phospholipid or cholesterol oxidation product.

8. (withdrawn) The method according to claim 6, wherein the detoxification enzyme is a cytochrome P450 enzyme or the catalase, a paraoxonase, a superoxide dismutase, a glutathione peroxidase, a

glutathione synthase, a glutathione reductase, a glutathione transferase, a glutamyl-cysteinyl synthase, a quinone reductase, a diaphorase, a thioredoxin, a glutaredoxin, a peroxiredoxin, an epoxide hydrolase, an aldehyde hydrolase, an aldo-keto reductase, a properdin, the selenoproteins P or W, an N-acetyl-transferase, a metallothionein, a sulfurtransferase, an alcohol dehydrogenase, an aldehyde dehydrogenase, a glutamate dehydrogenase, a dihydrodiol dehydrogenase, or a carboxyl esterase.

9. (withdrawn) The method according to claim 2, wherein the disorder comprises oxidative stress and testing for oxidative stress comprises determining the antioxidative capacity of HDL

10. (withdrawn) The method according to claim 4, wherein the reference value is selected from a reference range of 20 to 100 units per liter.

11. (withdrawn) The method according to claim 1, wherein the cardiovascular disease is coronary heart disease or cerebrovascular disease.

12. (withdrawn) The method according to claim 11, wherein the coronary heart disease is myocardial infarction.

13. (currently amended) Method of treatment of an individual to reduce ~~in order to protect the said individual against the risk of cardiovascular disease, the method comprising the steps of~~ which comprises testing the said individual for a disorder which ~~detrimentally affects~~ reduces the protective effect of HDL compared to the level of the protective effect of HDL in the absence of the disorder, identifying and selecting an individual the patient population that is free of the disorder ~~said condition~~, and treating the selected individual ~~in order to enhance~~ patient population to increase the HDL or HDL cholesterol level ~~of said individual~~, wherein said disorder is liver damage or oxidative stress.

14. (Cancelled)

15. (currently amended) The method according to claim ~~14~~ 13, wherein the disorder comprises liver damage and testing for liver damage comprises determining  $\gamma$ -glutamyltransferase activity or concentration in a serum or plasma sample and comparing it to a selected reference value for  $\gamma$ -glutamyltransferase.

16. (currently amended) The method according to claim ~~14~~ 13, wherein the disorder comprises liver damage and testing for liver damage comprises genotyping for mutations or polymorphisms that influence ~~influencing~~ serum or plasma  $\gamma$ -glutamyltransferase activity or concentration, ~~or~~ genotyping for mutations in the phase I and II enzymes, or analyzing RNA to measure the expression of these genes for  $\gamma$ -glutamyltransferase, phase I and/or phase II enzymes.

17. (currently amended) The method according to claim ~~14~~ 13, wherein the disorder comprises oxidative stress and testing for oxidative stress comprises determining serum or plasma paraoxonase activity or concentration.

18. (currently amended) The method according to claim ~~14~~ 13, wherein the disorder comprises oxidative stress and testing for oxidative stress comprises determining the antioxidative capacity of HDL.

19. (currently amended) The method according to claim 13, wherein the treatment to enhance the HDL or HDL cholesterol level is a drug treatment, ~~the drug being selected from the group~~

~~consisting of niacin, a statin, an apolipoprotein AI or AII synthesis enhancing agent, a PPAR alpha agonist such as a fibrate, a PPAR gamma or delta agonist, a sterol absorption inhibiting agent such as a resin, a CETP inhibitor, an ACAT inhibitor, a PLTP agonist, a LCAT agonist, a LPL agonist, a hepatic lipase agonist, a SR-B1 agonist, or a ABC1 (ATP binding cassette A1) agonist.~~

20. (original) The method according to claim 19, wherein the statin is selected from the group consisting of atorvastatin, fluvastatin, lovastatin, pravastatin and simvastatin.

21. (original) The method according to claim 19, wherein the fibrate is selected from the group consisting of bezafibrate, ciprofibrate, clofibrate, fenofibrate and gemfibrozil.

22. (original) The method according to claim 19, wherein the resin is selected from the group consisting of colestipol and cholestyramin.

23. (original) The method according to claim 13, wherein the treatment includes physical activity or physical exercise.

24. (withdrawn) The method according to claim 13, wherein the

treatment includes gene transfer and other kinds of gene therapy.

25. (withdrawn) A kit for identifying a condition in an individual in which condition an elevated serum or plasma HDL or HDL cholesterol concentration provides enhanced protection against cardiovascular disease, or for predicting an individual's response to HDL or HDL cholesterol elevating treatments, wherein the kit comprises means for testing the individual for a disorder which detrimentally affects the protective effect of HDL.

26. (withdrawn) The kit according to claim 25, wherein the additional condition is liver damage and the means comprise means for determining serum or plasma  $\gamma$ -glutamyltransferase or for genotyping genomic mutations and/or polymorphisms.

27. (withdrawn) The kit according to claim 25, wherein the additional condition is oxidative stress and the means comprise means for determining paraoxonase activity or concentration, the antioxidative capacity of HDL, or genotyping genomic mutations and polymorphisms.

28. (withdrawn) The kit according to claim 25 for assessing an individual's risk of cardiovascular disease further comprising

means for determining HDL or HDL cholesterol concentration in a serum or plasma sample of said individual.

29. (new) The method according to claim 19 wherein said drug is selected from the group consisting of niacin, a statin, an apolipoprotein I (AI) or apolipoprotein AII (AII) synthesis enhancing agent, a peroxisome proliferator-activated receptor (PPAR) alpha agonist, a PPAR gamma or delta agonist, a sterol absorption inhibiting agent, a cholesterol ester transfer protein (CETP) inhibitor, an acyl-coA cholesterol acyltransferase (ACAT) inhibitor, a phospholipid transfer protein (PLTP) agonist, a lecithin cholesterol acyltransferase (LCAT), agonist, a lipoprotein lipase (LPL) agonist, a hepatic lipase agonist, a scavenger receptor B1 (SR-B1) agonist, and a ATP-binding cassette A1 (ABC1) agonist.

30. (new) The method according to claim 29, wherein the PPAR alpha agonist a fibrate.

31. (new) The method according to claim 29, wherein the sterol absorption inhibiting agent is a resin.